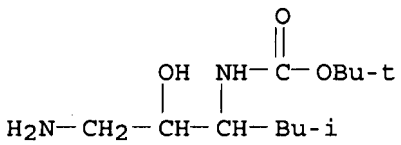
L4 ANSWER 190 OF 251 CAPLUS COPYRIGHT 2007 ACS on STN
AN 1989:595417 CAPLUS <<LOGINID::20070227>>
DN 111:195417
TI Preparation and testing of peptide amides as renin inhibitors
IN Luly, Jay R.; Dellaria, Joseph; Fung, Anthony K. L.; Kempf, Dale J.; Plattner, Jacob J.; Rosenberg, Saul H.; Sham, Hing L.
PA Abbott Laboratories, USA
SO U.S., 20 pp. Cont.-in-part of U.S. 4,645,759.
CODEN: USXXAM
DT Patent
LA English
FAN.CNT 4

| | PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|---|---|---|---|---|
| PI | US 4826815 | A | 19890502 | US 1987-946881 | 19870109 |
| | US 4645759 | A | 19870224 | US 1985-735491 | 19850517 |
| | JP 61033152 | A | 19860217 | JP 1985-134423 | 19850621 |
| | WO 8704349 | A1 | 19870730 | WO 1987-US54 | 19870116 |
| | W: JP | | | | |
| | RW: BE, CH, DE, FR, GB, IT, SE | | | | |
| | EP 258289 | A1 | 19880309 | EP 1987-900949 | 19870116 |
| | R: BE, CH, DE, FR, GB, IT, LI, SE | | | | |
| | JP 63503380 | T | 19881208 | JP 1987-500710 | 19870116 |
| PRAI | US 1984-623807 | A2 | 19840622 | | |
| | US 1985-735491 | A2 | 19850517 | | |
| | US 1986-820060 | A | 19860116 | | |
| | US 1986-820274 | A | 19860116 | | |
| | US 1986-850802 | A | 19860411 | | |
| | US 1986-862077 | A | 19860512 | | |
| | US 1987-946881 | A | 19870109 | | |
| | US 1987-946882 | A | 19870109 | | |
| | US 1987-946883 | A | 19870109 | | |
| | US 1987-946884 | A | 19870109 | | |
| | WO 1987-US54 | W | 19870116 | | |

OS CASREACT 111:195417; MARPAT 111:195417
AB ACHR1CR10R11WR2CHR3CONR4CHR5C(OH)R7CR8R9XR6 [I; A = H, C1-6 alkyl, aralkyl, HO, alkoxy, amino, R12COX1; R1 = C1-6 alkyl, PhCH2, β-naphthylmethyl, 4-MeOC6H4CH2; R2, R4, R7, R8, R9 = H, C1-6 alkyl; R3 = (OH-substituted) C1-6 alkyl, PhCH2, 4-HOC6H4CH2, 4-imidazolylmethyl; R6 = C1-6 alkyl, cycloalkyl, cycloalkylalkyl, aryl, alkylaryl, protecting group; R10, R11 = (H, OH), (H, H); or R10Rr11 = O; R12 = C1-6 alkyl, alkoxy, aralkoxy, amino, heterocyclylalkyl, (substituted) heterocyclyl; W = N, CH; X = NH, O, S, SO2, SO, CH2; X1 = NH, O, CH2, HNCH2], useful as antihypertensives, were prepared 3-Amino-2-hydroxy-5-methyl-1-phenylmercaptohexane (preparation given) in DMF was added to BOC-Phe-His-OH in DMF at -23° followed by hydroxybenzotriazole and DCC. After 2-5 h the mixture was kept at room temperature for 16 h to give the BOC-Phe-His amide of 3-amino-2-hydroxy-5-methyl-1-phenylmercaptohexane. The latter gave 56% inhibition of human renal renin at 10-6 M. The BOC-Phe-His amide of 3-amino-4-cyclohexyl-1-cyclohexylsulfonyl-2-hydroxybutane gave 81% inhibition at 10-8 M in the above screen. Approx. 50 I was prepared
IT 103127-80-6P
RL: SPN (Synthetic preparation); PREP (Preparation)
(preparation of, as intermediate for peptide renin inhibitor)
RN 103127-80-6 CAPLUS
CN Carbamic acid, [1-(2-amino-1-hydroxyethyl)-3-methylbutyl]-, 1,1-dimethylethyl ester, monohydrochloride (9CI) (CA INDEX NAME)



● HCl

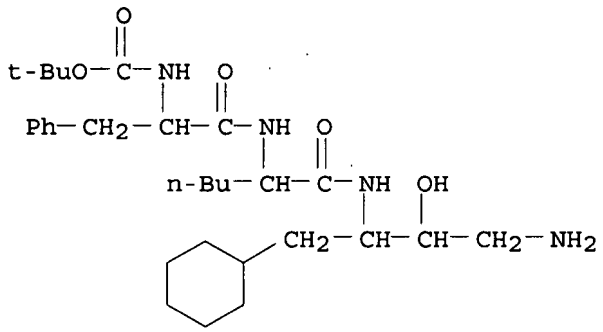
L4 ANSWER 191 OF 251 CAPLUS COPYRIGHT 2007 ACS on STN
AN 1989:497730 CAPLUS <<LOGINID::20070227>>
DN 111:97730
TI Preparation and testing of acylpeptide amides as cardiovascular agents and virucides
IN Weidmann, Beat
PA Sandoz-Patent-G.m.b.H., Fed. Rep. Ger.
SO Ger. Offen., 15 pp.
CODEN: GWXXBX
DT Patent
LA German
FAN.CNT 1

| | PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|---|---|---|---|---|
| PI | DE 3830825 | A1 | 19890323 | DE 1988-3830825 | 19880910 |
| | FR 2620451 | A1 | 19890317 | FR 1988-11952 | 19880912 |
| | FR 2620451 | B1 | 19931224 | | |
| | CH 677672 | A5 | 19910614 | CH 1988-3397 | 19880912 |
| | BE 1003071 | A5 | 19911112 | BE 1988-1045 | 19880912 |
| | GB 2209752 | A | 19890524 | GB 1988-21442 | 19880913 |
| | GB 2209752 | B | 19910605 | | |
| | JP 01151544 | A | 19890614 | JP 1988-231395 | 19880914 |
| | US 5045537 | A | 19910903 | US 1988-244220 | 19880914 |
| PRAI | DE 1987-3730895 | A1 | 19870915 | | |

AB R1O[(CH2)oOm(CH2)nWCOABCD [I; R1 = H, C1-20 alkyl, sugar residue, C2-30 alkylcarbonyl, C3-6 polyhydroxylalkylcarbonyl, phosphoryl, sulfo, aroyl, heteroaroyl, arylalkyl, biotinyl, D- or L-amino acid residue, etc.; W = O, CH2, imino; A, B, C = bond, NR2CHR3CO; D = NHCHR4CR5R6CR7R8XCONR9R10, NHCHR4CHR5CH2XSO2ZR9R10, NHCHR4CH2NHCHR11COR12; R2 = undefined; R3, R4 = hydrophilic or lipophilic amino acid side chain; R2R3 = (CH2)o; R5 = OH, amino; R6 = H; R5R6 = O; R7, R8 = F, H; R9, R10 = H, C1-5 alkyl, CHR11COR12; R11 = C1-5 alkyl, hydroxyalkyl; R12 = OH, C1-5 alkoxy, amino, alkylamino, aminomethylpyridyl, PhCH2, NH(CH2CH2O)mR1; X = O, NH, CR13R14; R13, R14 = H, F, R3; Z = N, CH; m = 1-20; n = 0-5; o = 2,3], useful as resin inhibitors, were prepared H-Thala-Nle-Chatin-Leu-α-Pic [Thala = (2S)-2-amino-3-(2-thienyl)propionyl, Chatin = (3S,4S)-4-amino-5-cyclohexyl-3-hydroxyvaleryl, α-Pic = 2-aminomethylpyridyl] in THF was treated with hydroxybenzotriazole, DCC, and 3,6,9,12-tetraoxatridecanoic acid (EG) in DMF to give EG-Thala-Nle-Chatin-Leu-α-Pic. I inhibited human plasma renin at 10-5-10-11 M. I also completely eliminated feline leukemia virus in cats after 14 days.

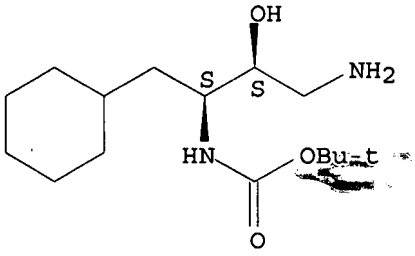
IT 118546-39-7
RL: RCT (Reactant); RACT (Reactant or reagent)
(acylation of, by iso-Pr isocyanate, in preparation of cardiovascular agent and virucide)

RN 118546-39-7 CAPLUS
CN L-Norleucinamide, N-[(1,1-dimethylethoxy)carbonyl]-L-phenylalanyl-N-[3-amino-1-(cyclohexylmethyl)-2-hydroxypropyl]-, [S-(R*,R*)]- (9CI) (CA INDEX NAME)

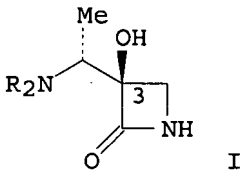


IT 108868-90-2
RL: RCT (Reactant); RACT (Reactant or reagent)
(reaction of, in preparation of cardiovascular agent and virucide)
RN 108868-90-2 CAPLUS
CN Carbamic acid, [3-amino-1-(cyclohexylmethyl)-2-hydroxypropyl]-, 1,1-dimethylethyl ester, [S-(R*,R*)]- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



L4 ANSWER 197 OF 251 CAPLUS COPYRIGHT 2007 ACS on STN
AN 1989:135696 CAPLUS <<LOGINID::20070227>>
DN 110:135696
TI Synthesis of an analog of tabtoxinine as a potential inhibitor of D-alanine:D-alanine ligase (ADP forming)
AU Greenlee, William J.; Springer, James P.; Patchett, Arthur A.
CS Merck Sharp and Dohme Res. Lab., Rahway, NJ, 07065, USA
SO Journal of Medicinal Chemistry (1989), 32(1), 165-70
CODEN: JMCMAR; ISSN: 0022-2623
DT Journal
LA English
OS CASREACT 110:135696
GI



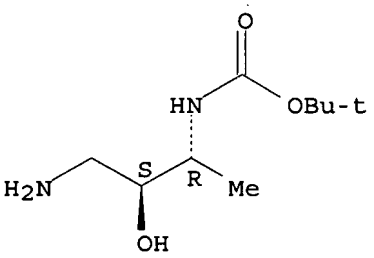
I

AB The design and synthesis of a potential inhibitor of D-alanine:D-alanine ligase (ADP forming) (EC 6.3.2.4) are described. This enzyme, which catalyzes the second step in the biosynthesis of bacterial peptidoglycan, is believed to generate D-alanylphosphate as an enzyme-bound intermediate. With tabtoxinine (a potent inhibitor of glutamine synthetase) as a model, β-lactams (3R)- and (3S)-I (R = H) were synthesized as potential precursors of a D-alanylphosphate mimic. The structure of I (R = CH2CH:CH2) was proved by x-ray crystallog.
IT 119391-97-8P 119413-59-1P
RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)
(preparation and cyclocondensation reaction of, with phthalic anhydride)
RN 119391-97-8 CAPLUS
CN Carbamic acid, (3-amino-2-hydroxy-1-methylpropyl)-, 1,1-dimethylethyl ester, [S-(R*,S*)]-, monoacetate (salt) (9CI) (CA INDEX NAME)

CM 1

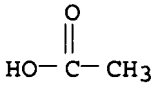
CRN 119391-96-7
CMF C9 H20 N2 O3

Absolute stereochemistry.



CM 2

CRN 64-19-7
CMF C2 H4 O2

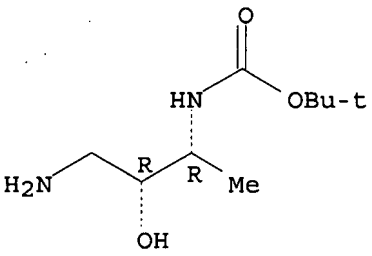


RN 119413-59-1 CAPLUS
CN Carbamic acid, (3-amino-2-hydroxy-1-methylpropyl)-, 1,1-dimethylethyl ester, [R-(R*,R*)]-, monoacetate (salt) (9CI) (CA INDEX NAME)

CM 1

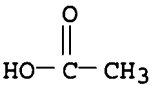
CRN 119413-58-0
CMF C9 H20 N2 O3

Absolute stereochemistry.



CM 2

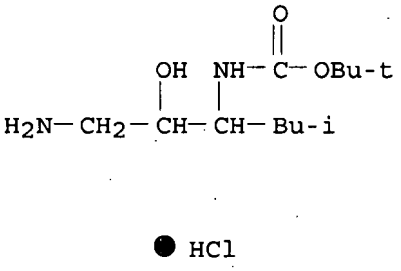
CRN 64-19-7
CMF C2 H4 O2



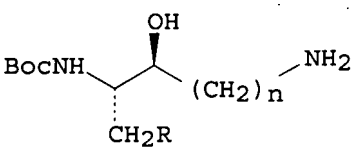
L4 ANSWER 208 OF 251 CAPLUS COPYRIGHT 2007 ACS on STN
AN 1986:443336 CAPLUS <<LOGINID::20070227>>
DN 105:43336
TI Renin inhibiting compounds
IN Luly, Jay Richard; Dellaria, Joseph F., Jr.; Plattner, John Jacob
PA Abbott Laboratories, USA
SO Eur. Pat. Appl., 35 pp.
CODEN: EPXXDW
DT Patent
LA English
FAN.CNT 4

| | PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|---|---|---|---|---|
| PI | EP 172347 | A2 | 19860226 | EP 1985-107375 | 19850618 |
| | EP 172347 | A3 | 19890405 | | |
| | R: BE, CH, DE, FR, GB, IT, LI, NL, SE | | | | |
| | US 4645759 | A | 19870224 | US 1985-735491 | 19850517 |
| | JP 61033152 | A | 19860217 | JP 1985-134423 | 19850621 |
| PRAI | US 1984-623807 | A | 19840622 | | |
| | US 1985-735491 | A | 19850517 | | |

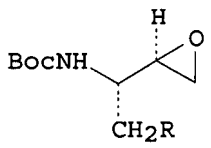
OS CASREACT 105:43336; MARPAT 105:43336
AB Title compds. RnZ1CHR1CONR2CHR3CONR4CHR5CR7(OH)CR8R9Z2R6 (n = 0,1; R = N-protecting group; Z1 = H, OH, alkyl, arylalkyl, NH; R1, R3 and R5 are alkyl, amino acid side chains; R2 R4, R7, R8, and R9 are H, alkyl; Z2 = NH, O, S, SO, SO2; R6 = alkyl, cycloalkyl, cycloalkylalkyl, aryl, etc.), which showed antihypertensive activity, were prepared A protected histidine was amidated, and the product was deprotected and coupled to give BOC-Phe-His-NHCH(CH2CHMe2)CH(OH)CH2S(CH2)3Ph.
IT 103127-80-6P
RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)
(preparation and reaction of)
RN 103127-80-6 CAPLUS
CN Carbamic acid, [1-(2-amino-1-hydroxyethyl)-3-methylbutyl]-, 1,1-dimethylethyl ester, monohydrochloride (9CI) (CA INDEX NAME)



L4 ANSWER 205 OF 251 CAPLUS COPYRIGHT 2007 ACS on STN
AN 1987:459446 CAPLUS <<LOGINID::20070227>>
DN 107:59446
TI Novel renin inhibitors containing analogs of statine retro-inverted at the C-termini. Specificity at the P2 histidine site
AU Rosenberg, Saul H.; Plattner, Jacob J.; Woods, Keith W.; Stein, Herman H.; Marcotte, Patrick A.; Cohen, Jerome; Perun, Thomas J.
CS Cardiovasc. Res. Div., Abbott Lab., Abbott Park, IL, 60064, USA
SO Journal of Medicinal Chemistry (1987), 30(7), 1224-8
CODEN: JMCMAR; ISSN: 0022-2623
DT Journal
LA English
OS CASREACT 107:59446
GI



I



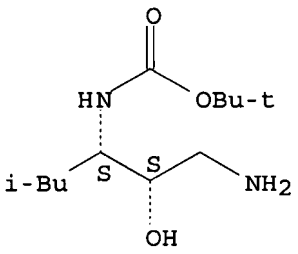
II

AB Substituted 1,3- and 1,4-diamines I (Boc = Me3CO2C; R = CHMe2, n = 1; R = cyclohexyl, n = 1, 2) were prepared from epoxides II. These diamines were incorporated into renin inhibitors (IC50 = 4-1500 nM) replacing the Leu-Val scissile bond in small peptide analogs of angiotensinogen. Replacement of the P2 histidine imidazole with other heterocycles maintained or enhanced binding while changing the overall basicity of the inhibitor. Substitution of O-methyltyrosine for the P3 phenylalanine suppressed chymotrypsin cleavage of the P3-P2 bond.
IT 108868-53-7P 108868-91-3P
RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)
(preparation and N-acylation of)
RN 108868-53-7 CAPLUS
CN Carbamic acid, [1-(2-amino-1-hydroxyethyl)-3-methylbutyl]-, 1,1-dimethylethyl ester, [S-(R*,R*)]-, monoacetate (salt) (9CI) (CA INDEX NAME)

CM 1

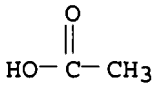
CRN 108868-52-6
CMF C12 H26 N2 O3

Absolute stereochemistry.



CM 2

CRN 64-19-7
CMF C2 H4 O2

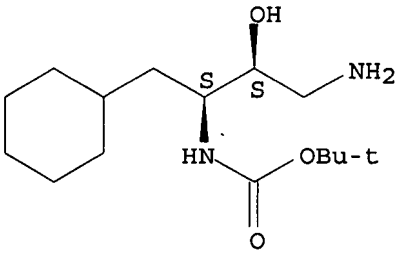


RN 108868-91-3 CAPLUS
CN Carbamic acid, [3-amino-1-(cyclohexylmethyl)-2-hydroxypropyl]-, 1,1-dimethylethyl ester, [S-(R*,R*)]-, monoacetate (salt) (9CI) (CA INDEX NAME)

CM 1

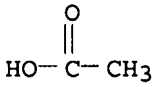
CRN 108868-90-2
CMF C15 H30 N2 O3

Absolute stereochemistry.



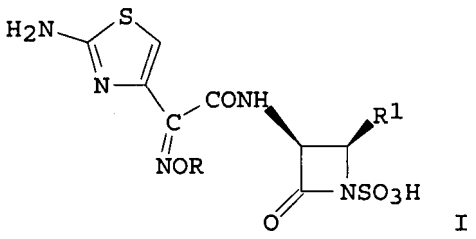
CM 2

CRN 64-19-7
CMF C2 H4 O2



L4 ANSWER 206 OF 251 CAPLUS COPYRIGHT 2007 ACS on STN
AN 1986:552768 CAPLUS <<LOGINID::20070227>>
DN 105:152768
TI Synthesis of carumonam (AMA-1080) and a related compound starting from (2R,3R)-epoxysuccinic acid
AU Sendai, Michiyuki; Hashiguchi, Shohei; Tomimoto, Mitsumi; Kishimoto, Shoji; Matsuo, Taisuke; Ochiai, Michihiko
CS Cent. Res. Div., Takeda Chem. Ind., Ltd., Osaka, 532, Japan
SO Chemical & Pharmaceutical Bulletin (1985), 33(9), 3798-810
CODEN: CPBTAL; ISSN: 0009-2363
DT Journal
LA English
OS CASREACT 105:152768

GI



I

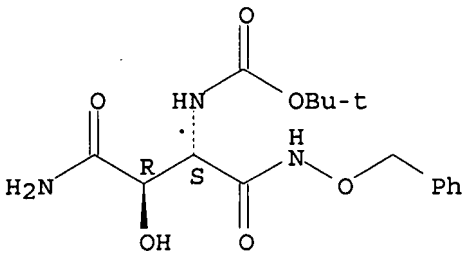
AB Several 4-carbamoyl-2-azetidinone-1-sulfonic acid derivs. I [R = Me, CH2CO2H, CMe2CO2H; R1 = CH2OCONH2, CONR2R3 (R2 = H, Me; R3 = H, Me)] were prepared to improve the antibacterial activity of sulfazecin. I (R = CMe2CO2H, R1 = CONH2) showed potent antibacterial activity, comparable to that of carumonam against gram-neg. bacteria. Efficient synthetic pathways to prepare I (R = CH2CO2H, R1 = CH2O2CNH2; R = CMe2CO2H, R1 = CONH2) in large quantities were developed based on (2R,3R)-epoxysuccinic acid, an easily accessible fermentation product.

IT 98377-00-5P
RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)
(preparation and mesylation of)

RN 98377-00-5 CAPLUS

CN Carbamic acid, [3-amino-2-hydroxy-3-oxo-1-[[(phenylmethoxy)amino]carbonyl]propyl]-, 1,1-dimethylethyl ester, [R-(R*,S*)]- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

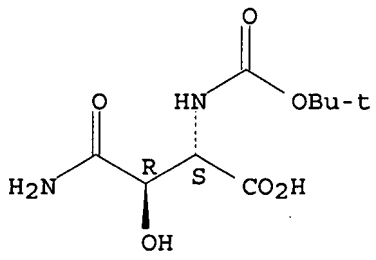


IT 98463-52-6P
RL: SPN (Synthetic preparation); PREP (Preparation)
(preparation and protection of carboxylic group)

RN 98463-52-6 CAPLUS

CN L-Asparagine, N2-[(1,1-dimethylethoxy)carbonyl]-3-hydroxy-, erythro- (9CI) (CA INDEX NAME)

Absolute stereochemistry.


O
HN
OBu-t
O
R
S
H2N
CO2H
OH

L4 ANSWER 1 OF 1 REGISTRY COPYRIGHT 2007 ACS on STN
RN 119391-96-7 REGISTRY
CN Carbamic acid, (3-amino-2-hydroxy-1-methylpropyl)-, 1,1-dimethylethyl ester, [S-(R*,S*)]- (9CI) (CA INDEX NAME)
FS STEREOSEARCH
MF C9 H20 N2 O3
CI COM
SR CA
LC STN Files: CHEMCATS

Absolute stereochemistry.